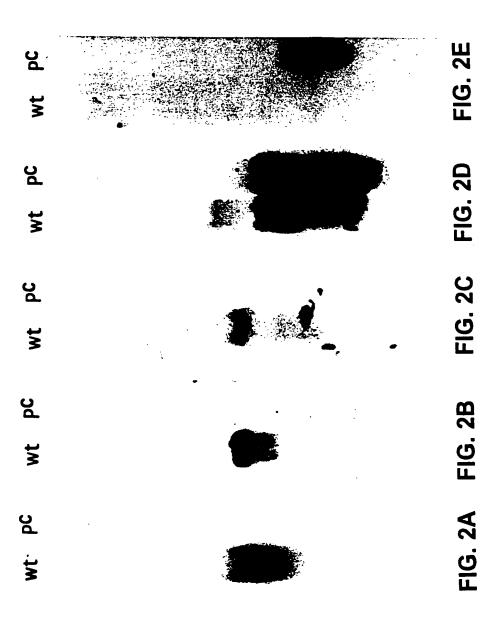


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Applicant(s): Samuel I. Miller
SALMONELLA SECRETED PROTEINS AND USES THEREOF







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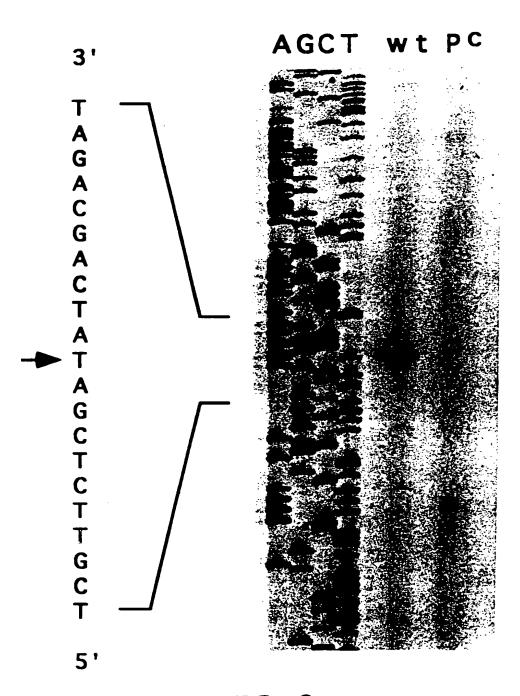


FIG. 3





|      |     | * ** +* ** * * * * ** ** ** +* +                     |            |        |
|------|-----|--|------------|--------|
| PrgI | 1   | MATPWSGYLDDVSAKFDTGVDNLQTQVTEALDKLAAKPSDP            | 41         |        |
| MxiH | ī   | MSVTVPNDDWTLSSLSETFDDGTQTLQGELTLALDKLAKNPSNP         | 44         |        |
| YscF | ī   | MSNFSGFTKGNDIADLDAVAQTLKKPADDANKAVNDSIAALKDTPDNP     | 48         |        |
|      | _   |  |            |        |
|      |     | *+++ *+***** ***+**+** ++ ***+*                      |            |        |
| PrgI | 42  | ALLAAYQSKLSEYNLYRNAQSNTVKVFKDIDAAIIQNFR 80 SEQ ID NO | ):32:      |        |
| MxiH | 45  | QLLAEYQSKLSEYTLYRNAQSNTVKVIKDVDAAIIQNFR 83 SEQ ID NO | ):33:      |        |
| YscF | 49  | ALLADLQHSINKWSVIYNISSTIVRSMKDLMQGILQKFP 87 SEQ ID NO | 0:34:      |        |
|      |     |  |            |        |
|      |     | FIG. 4A  |            |        |
|      |     |  |            |        |
|      |     |  |            |        |
|      |     | +++++ + +++++ + +                                    |            |        |
| PrgJ | 1   | MSIATIVPENAV. IGQAVNIRSMETDIVSLDDRLLQAFSGSAIATAVDKQT | 50         |        |
| MxiI | 1   | MNYIYPVNQVDIIKASDFQSQEISSLEDVVSAKYSDIKMDTDIQVSQ      | 47         |        |
|      |     |  |            |        |
|      |     | +              |            | SEQ ID |
| PrgJ | 51  | ITNRIEDPNLVTDPKELAISQEMISDYNLYVSMVSTLTRKGVGAVETLLRS  |            | NO:35  |
| MxiI | 48  | IMEMVSNPESL.NPESLAKLQTTLSNYSIGVSLAGTLARKTVSAVETLLKS  | 97         | SEQ ID |
|      |     | FIG. 4D  |            | NO:36  |
|      |     | FIG. 4B  |            |        |
|      |     |  |            |        |
|      |     |  |            |        |
|      |     | *** * * +***+ ++* *+ ** + ***+ *** + +               |            |        |
| PrqK | 1   | MIRRYLYTELLYMTLAGCKDK.DLLKGLDQEQANEVIAVLQMHNIEANKI   | 49         |        |
| MxiJ | ī   | MIRYKGFILFILLMLIGCEOREELISNLSQRQANEIISVLERHNITARKV   | 50         |        |
| YscJ | ī   | MKVKTSLSTLILILFLTGCKVDLYTGISQKEGNEMLALLRQEGLSADKE    | 49         |        |
|      | _   |  |            |        |
|      |     | * ** * * + * ** +* * + ** * * **++ +*+*++*           |            |        |
| PrgK | 50  | DSGKLGYSITVAEPDFTAAVYWIKTYQLPPRPRVEIAQMFPADSLVSSPR   | 99         |        |
| MxiJ | 51  | DGGKOGTSVOVEKGTFASAVDLMRMYDLPNPERVDISQMFPTDSLVSSPR   | 100        |        |
| YscJ | 50  | PDKDGKIKLLVEESDVAQAIDILKRKGYPHESFSTLQDVFPKDGLISSPI   | 9 <b>9</b> |        |
|      |     |  |            |        |
|      |     | *+*+++**+**  |            |        |
| PrqK | 100 | AEKARLYSAIEQRLEQSLQTMEGVLSARVHISYDIDAGENGRPPKPVHLS   | 149        |        |
| MxĺJ |     | AFKARLYSATEORLEOSLVSIGGVISAKIHVSYDLEEKNISSKPMHIS     | 148        |        |
| YscJ |     | EELARLNYAKAQEISRTLSEIDGVLVARVHVVLPEEQNNKGKKGVAASAS   | 149        |        |
|      |     |  |            |        |
|      |     | * * * ++****** ** +**++ * *                          |            |        |
| PraK | 150 | ALAVYERGSPLAHQISDIKRFLKNSFADVDYDNISVVLSERSDAQL       | 195        |        |
| MxiJ | 149 | VTATVDSPKESELLVSNIKRFLKNTFSDVKYENISVILTPKEEYVY       | 195        |        |
| YscJ | 150 | VFIKHAADIQFDTYIPQIKQLVNNSIEGLAYDRISVILVPSVDVRQSSHL   | 199        |        |
|      |     |  |            |        |
| •    |     | * ** * * *** +* * +                                  |            |        |
| PrqK | 196 | QAPGTPVKRNSFATSWIVLIILLSVMSAGFGVWYYKNHYARNKKGITA     | 243        | SEQ ID |
| MxiJ |     | TNVOPVKEVKSEFLTNEVIYLFLGMAVLVVILLVWAFKTGWFKRNKI      | 270        | NO:3/  |
| YscJ |     | P. RNTSILSIQVSEESKGRLIGLLSLLILLLPVTNLAQYFWLQRKK      | 244        | SEQ ID |
|      |     |  |            | NO:38  |
| PrqK | 243 | DDKAKSSNE 252 SEQ ID NO:39                           |            |        |
|      |     |  |            |        |



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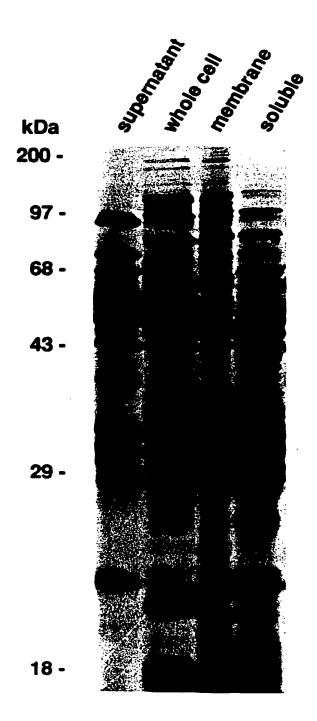


FIG. 5



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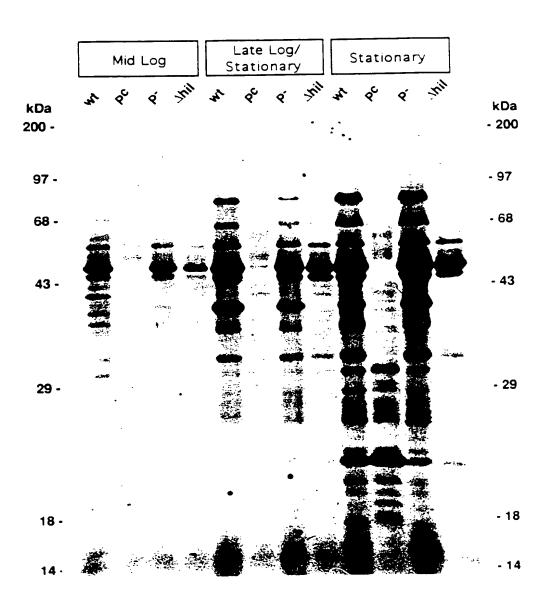


FIG. 6

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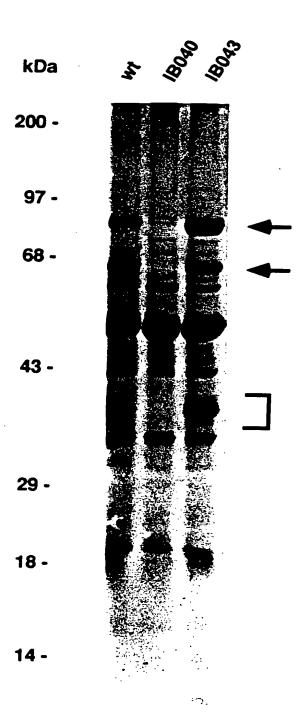


FIG. 7



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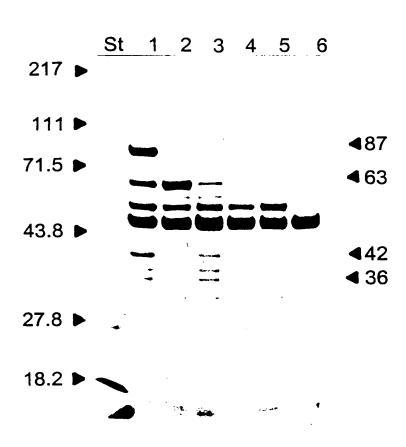


FIG. 8

<u>- M</u>

<u>(P</u>

P lac

EE638 [pCH004]

EE638 [pCH002]

Appln No.: 09/068,804 Applicant(s): Samuel I. Miller

1kb

B

lacZY4

E(N) (N)

മ

E(N) (N)

**EE633** 

lacZY11-6

prgH

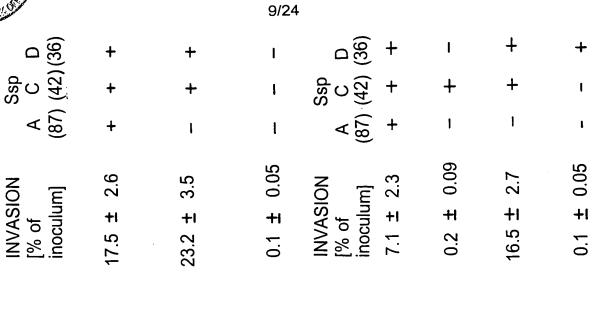
sspA\*

sspB\* sspC sspD

spaT

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**-** 🗠

 $\mathbf{\omega}$ 

E(N) (N)

**EE638** 

FIG. 10

P<u>lac</u>

EE638 [pCH006]

EE638 [pCH005]





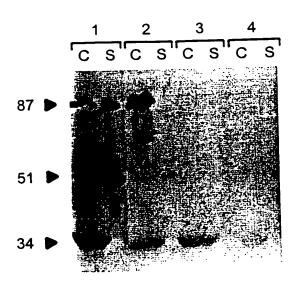


FIG. 11





RKAEETNRIMGCIGKVIGALLTIVSVVAAVFTGGASLALAAVGLAVMVADEIVKAATGVSFIQQALNPIMEHVLKPLMELIGKAITKALEGIGVDKKTAE





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1 MINIQNYSASPHPGIVAERPQTPSASEHVETAVVPSTTEHRGTDIISLSQAATKIHQAQQTLQ...STPPISEENNDERTLARQQLTSSLNALAKSGVSL MNITTLINSISTSSFSPNNTNGSSTETVNSDIKTTTSSHPVSSLTMLNDTLHNIRTTNQALKKELSQKTLTKTSLEETALHSSQISMDVNKSAQLLDIL SAEQ...NENLRSAF.SAPTSALFSASPWAQPRTTISDAEIWDMVSQNISAIGDSYLGVYENVVAVYTDFYQAFSDILSKMGGWILP.GKDGNTVKLDVT IPAD 100 SRNEYPINKDARELLHSAPKEAELDGDQM.....ISHRELMAKIANSINDINEQYLKVYEHAVSSYTQMYQDFSAVLSSLAGWISPGGNDGNSVKLQVN SLKNDI.NSLVNKYNQI NSNTVLFPAQSGSGVKVATEAEARQMLSELNLPNSCLKSYGSGYVVTVDLTPLQKMVQDI DGI.GAPGKDSKLEMDNAKYQAWQS IPAD 194 SLKKALEELKEKYK....DKPLYPANNT....VSQEQANKWLTELGGTIGKVSQKNGGYVVSINMTPIDNMLKSLDNLGGNG...EVVLDNAKYQAWNA IPAD 282 GFSAEDETWKNNLQTLVQKYSNANSIFDNLVKVLSSTISSCTDTDKLFLHF 332 SEQ ID NO:45 SSPD 293 GFKAQEENMKTTLQTLTQKYSNANSLYDNLVKVLSSTISSSLETAKSFLQG 343 SEQ ID NO:44 86 SspD 193 SspD SspD IpaD

SEQ ID NO:46 1 MVTSVRTQPPVIMPGMQTEIKTQATNLAANLSAVRESATATLSGEIKGPQLEDFPALIKQASLD 64

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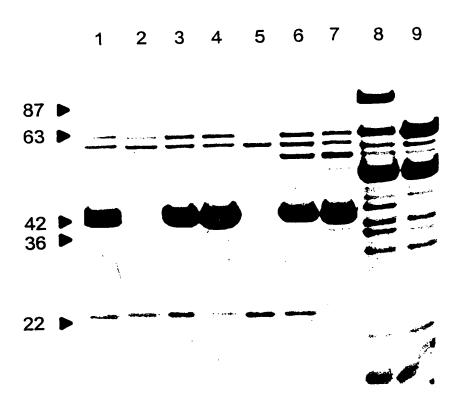


FIG. 17

40

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6

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 $\geq$ 

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spa15

S

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 $\geq$ 

ipaB

S. flexneri

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15/24 orgA Þ 0 ပ **SSPB** ORF1 prgH S hilA  $\alpha$ **MXIH** Q ٥. Ш ipgD icsB ipgA S  $\boldsymbol{\omega}$ Θ  $\circ$ P

Ш

invH invF

S. typhimurium

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#### SEQ ID NO:1:

| CGCAAAGCCG | AGGAAACGAA | CCGCATTATG | GGATGTATCG | GGAAAGTCCT | CGGCGCGCTG        | 60  |
|------------|------------|------------|------------|------------|-------------------|-----|
| CTAACCATTG | TCAGCGTTGT | GGCCGCTGTT | TTTACCGGTG | GGGCGAGTCT | GGCGCTGGCT        | 120 |
| GCGGTGGGAC | TTGCGGTAAT | GGTGGCCGAT | GAAATTGTGA | AGGCGGCGAC | GGGAGTGTCG        | 180 |
| TTTATTCAGC | AGGCGCTAAA | CCCGATTATG | GAGCATGTGC | TGAAGCCGTT | AATGGAGCTG        | 240 |
| ATTGGCAAGG | CGATTACCAA | AGCGCTGGAA | GGATTAGGCG | TCGATAAGAA | AACGGCAGAG        | 300 |
| ATGGCCGGCA | GCATTGTTGG | TGCGATTGTC | GCCGCTATTG | CCATGGTGGC | GGTCATTGTG        | 360 |
| GTGGTCGCAG | TTGTCGGGAA | AGGCGCGGCG | GCGAAACTGG | GTAACGCGCT | GAGCAAAATG        | 420 |
| ATGGGCGAAA | CGATTAAGAA | GTTGGTGCCT | AACGTGCTGA | AACAGTTGGC | GCAAAACGGC        | 480 |
| AGCAAACTCT | TTACCCAGGG | GATGCAACGT | ATTACTAGCG | GTCTGGGTAA | TGTGGGTAGC        | 540 |
| AAGATGGGCC | TGCAAACGAA | TGCCTTAAGT | AAAGAGCTGG | TAGGTAATAC | CCTAAATAAA        | 600 |
| GTGGCGTTGG | GCATGGAAGT | CACGAATACC | GCAGCCCAGT | CAGCCGGTGG | TGTTGCCGAG        | 660 |
| GGCGTATTTA | TTAAAAATGC | CAGCGAGGCG | CTTGCTGATT | TTATGCTCGC | CCGTTTTGCC        | 720 |
| ATGGATCAGA | TTCAGCAGTG | GCTTAAACAA | TCCGTAGAAA | TATTTGGTGA | <u>AAACCAGAAG</u> | 780 |
| GTAACGGCGG | AACTGCAAAA | AGCCATGTCT | TCTGCGGTAC | AGCAAAATGC | GGATGCTTCG        | 840 |
| CGTTTTATTC | TGCGCCAGAG | TCGCGCATAA |            |            |                   | 870 |
|            |            |            |            |            |                   |     |

## FIG. 19

### SEQ ID NO:2:

| ATGTTAATTA GTAATGTGGG | AATAAATCCC | GCCGCTTATT | TAAATAATCA | TTCTGTTGAG | 60   |
|-----------------------|------------|------------|------------|------------|------|
| AATAGTTCAC AGACAGCTTC | GCAATCCGTT | AGCGCTAAAG | ATATTCTGAA | TAGTATTGGT | 120  |
| ATTAGCAGCA GTAAAGTCAG | TGACCTGGGG | TTGAGTCCTA | CACTGAGCGC | GCCTGCGCCA | 180  |
| GGGGTATTAA CGCAAACCCC | CGGAACGATC | ACGTCCTTTT | TAAAAGCCAG | TATTCAAAAT | 240  |
| ACCGACATGA ATCAGGATTT | GAATGCTCTG | GCAAATAATG | TCACGACTAA | AGCGAATGAG | 300  |
| GTTGTGCAAA CCCAGTTACG | CGAGCAGCAG | GCAGAAGTCG | GAAAGTTTTT | TGATATTAGC | 360  |
| GGAATGTCTT CCAGTGCCGT | TGCGCTGTTG | GCTGCCGCGA | ATACGTTAAT | GCTGACGTTG | 420  |
| AACCAGGCTG ATAGCAAACT | GTCTGGTAAG | TTGTCATTAG | TCAGTTTTGA | TGCAGCTAAA | 480  |
| ACGACGGCAA GCTCCATGAT | GCGCGAAGGG | ATGAATGCGT | TGTCCGGTAG | TATTTCCCAG | 540  |
| AGCGCGCTTC AGTTGGGGAT | CACTGGCGTG | GGCGCCAAAC | TGGAATATAA | GGGGCTGCAG | 600  |
| AATGAAAGAG GCGCGCTTAA | ACATAATGCC | GCGAAGATCG | ATAAACTGAC | CACTGAAAGC | 660  |
| CACAGTATTA AAAACGTGCT | GAACGGGCAG | AATAGCGTCA | AACTCGGTGC | TGAAGGCGTC | 720  |
| GATTCTCTGA AATCGTTAAA | TATGAAGAAA | ACCGGTACCG | ATGCGACGAA | AAATCTTAAT | 780  |
| GATGCGACGC TTAAATCTAA | TGCCGGAACC | AGCGCCACGG | AAAGTCTGGG | TATTAAAGAC | 840  |
| AGTAATAAAC AAATCTCCCC | TGAACATCAG | GCTATTCTGT | CGAAACGTCT | TGAGTCTGTC | 900  |
| GAATCCGATA TTCGTCTTGA | GCAGAATACC | ATGGATATGA | CCCGAATCGA | TGCGCGCAAG | 960  |
| ATGCAGATGA CGGGCGATCT | GATTATGAAG | AACTCGGTCA | CGGTCGGTGG | TATTGCAGGG | 1020 |
| GCGTCCGGGC AGTACGCCGC | TACTCAGGAA | CGTTCCGAGC | AGCAAATTAG | CCAGGTGAAT | 1080 |
| AACCGGGTTG CCAGCACCGC | ATCGGACGAA | GCCCGTGAAA | GTTCACGTAA | ATCGACCAGC | 1140 |
| CTGATTCAGG AAATGCTGAA | AACAATGGAG | AGCATTAACC | AGTCGAAAGC | ATCCGCACTC | 1200 |
| GCTGCTATCG CAGGCAATAT | TCGCGCTTAA |            |            |            | 1230 |
|                       |            | -          |            |            |      |



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### SEQ ID NO:3:

| ATGCTTAATA | TTCAAAATTA | TTCCGCTTCT  | CCTCATCCGG | GGATCGTTGC | CGAACGGCCG        | 60   |
|------------|------------|-------------|------------|------------|-------------------|------|
| CAGACTCCCT | CGGCGAGCGA | GCACGTCGAG  | ACTGCCGTGG | TACCGTCTAC | CACAGAACAT        | 120  |
| CGCGGTACAG | ATATCATTTC | ATTATCGCAG  | GCGGCTACTA | AAATCCACCA | GGCACAGCAG        | 180  |
| ACGCTGCAGT | CAACGCCACC | GATCTCTGAA  | GAGAATAATG | ACGAGCGCAC | GCTGGCGCGC        | 240  |
| CAGCAGTTGA | CCAGCAGCCT | GAATGCGCTG  | GCGAAGTCCG | GCGTGTCATT | ATCCGCAGAA        | 300  |
| CAAAATGAGA | ACCTGCGGAG | CGCGTTTTCT  | GCGCCGACGT | CGGCCTTATT | TAGCGCTTCG        | 360  |
| <u> </u>   | AGCCGAGAAC | AACCATTTCT  | GATGCTGAGA | TTTGGGATAT | GGTTTCCCAA        | 420  |
| CCTATGGCGC | CGATAGGTGA | CAGCTATCTG  | GGCGTTTATG | AAAACGTTGT | CGCAGTCTAT        | 480  |
| AATATATCGG |            | CAGTGATATT  | CTTTCCAAAA | TGGGAGGCTG | GTTATTACCA        | 540  |
| ACCGATTTT  | ATCAGGCCTT | <del></del> |            |            | TTTAAACAGT        | 600  |
| GGTAAGGACG | GTAATACCGT | TAAGCTAGAT  | GTTACCTCAC | TCAAAAATGA |                   |      |
| TTAGTCAATA | AATATAATCA | AATAAACAGT  | AATACCGTTT | TATTTCCAGC | GCAGTCAGGC        | 660  |
| AGCGGCGTTA | AAGTAGCCAC | TGAAGCGGAA  | GCGAGACAGT | GGCTCAGTGA | <u>ATTGAATTTA</u> | 720  |
| CCGAATAGCT | GCCTGAAATC | TTATGGATCC  | GGTTATGTCG | TCACCGTTGA | TCTGACGCCA        | 780  |
|            | TGGTTCAGGA | TATTGATGGT  | TTAGGCGCGC | CGGGAAAAGA | CTCAAAACTC        | 840  |
| TTACAAAAAA |            |             |            | TTAAAGCGCA | GGAAGAAAAT        | 900  |
| GAAATGGATA | ACGCCAAATA | TCAAGCCTGG  |            |            |                   | 960  |
| ATGAAAACCA | CATTACAGAC | GCTGACGCAA  | AAATATAGCA | ATGCCAATTC | ATTGTACGAC        |      |
| AACCTGGTAA | AAGTGCTGAG | CAGTACGATA  | AGTAGCAGCC | TGGAAACCGC | CAAAAGCTTC        | 1020 |
|            |            |             |            |            |                   | 1032 |
| CTGCAAGGAT | <u>AA</u>  |             |            |            |                   |      |



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ATGGTTACAA GTGTAAGGAC TCAGCCCCC GTCATAATGC CAGGTATGCA GACCGAGATC

AAAACGCAGG CCACGAATCT TGCGGCGAAT CTTTCCGCAG TCAGAGAAAG TGCCACAGCG

ACGCTGTCAG GGGAAATTAA AGGCCCGCAA CTGGAAGATT TTCCCGCGCT GATCAAACAG

GCGAGTCTGG ATGC SEQ ID NO:4

RKAEETNRIMGCIGKVLGALLTIVSVVAAVFTGGASLALAAVGLAVMVADEIVKAATGVS FIQQALNPIMEHVLKPLMELIGKAITKALEGLGVDRKRQRWPAALLVRLSPLCHGDAVIV VVAVVGKGAAAKLGNALSKMMGETIKKLVPNVLKQLAQNGSKLFTQGMQRITSGLGNVGS KMGLQTNALSKELVGNTLNKVALGMEVTNTAAQSAGGVAEGVFIKNASEALADFMLARFA MDQIQQWLKQSVEIFGENQKVTAELQKAMSSAVQQNADASRFILRQSRAZ SEQ ID NO:5

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MLISNVGINPAAYLNNHSVENSSQTASQSVSAKDILNSIGISSSKVSDLGLSPTLSAPAP GVLTQTPGTITSFLKASIQNTDMNQDLNALANNVTTKANEVVQTQLREQQAEVGKFFDIS GMSSSAVALLAAANTLMLTLNQADSKLSGKLSLVSFDAAKTTASSMMREGMNALSGSISQ SALQLGITGVGAKLEYKGLQNERGALKHNAAKIDKLTTESHSIKNVLNGQNSVKLGAEGV DSLKSLNMKKTGTDATKNLNDATLKSNAGTSATESLGIKDSNKQISPEHQAILSKRLESV ESDIRLEQNTMDMTRIDARKMQMTGDLIMKNSVTVGGIAGASGQYAATQERSEQQISQVN NRVASTASDEARESSRKSTSLIQEMLKTMESINQSKASAL AAIAGNIRAZ SEQ ID NO:6

MLNIQNYSASPHPGIVAERPQTPSASEHVETAVVPSTTEHRGTDIISLSQAATKIHQAQQ TLQSTPPISEENNDERTLARQQLTSSLNALAKSGVSLSAEQNENLRSAFSAPTSALFSAS PMAQPRTTISDAEIWDMVSQNISAIGDSYLGVYENVVAVYTDFYQAFSDILSKMGGWLLP GKDGNTVKLDVTSLKNDLNSLVNKYNQINSNTVLFPAQSGSGVKVATEAEARQWLSELNL PNSCLKSYGSGYVVTVDLTPLQKMVQDIDGLGAPGKDSKLEMDNAKYQAWQSGFKAQEEN MKTTLQTLTQKYSNANSLYDNLVKVLSSTISSSLETAKSFLQGZ SEQ ID NO:7

MVTSVRTQPPVIMPGMQTEIKTQATNLAANLSAVRESATATLSGEIKGPQLEDFPALIKQ ASLD SEQ ID NO:8



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ATCAGCTTGCCGTCGTCATAAGCAACTGGGCTTGCATTGCTTTTAGTTGTACAAACTGTGCAGGC
GTCTTCCAGCATTCTATTGTTCCGTGAATCCGGAAATCTGCACGTACCTGCTCCAGATTACTATG
AGGATTATCCTTAAGTACAAGGGCCGCCGCCATCGTTCCGGTTCTTCCCACTCCGCCCAGACAAT
GAATCATCGGTAAATGCTTATCTGATGAACTACGCCCCGGCGCGCCCATTTTGGTTACTATTTTC
ACCCTATCCGCCAGGTATTCTAACTGATCCGTAGACGGTAACGGCTGGTGATCTGGCCAATTTTT
CACATGCAATACCGGGATTGTATACCGCTTTCCCCGCAGGACAGTTGCATATTGTATTGGTCTAT
CGCTTCTCCCTGACTGGCTGAGCTCTCTTTTTTGGCTGTTTGTATCCACCTCGCCAAAGGTGTAGC
TCCCTCTGAAATAGGTGGTAATTGTTTTTGCCTGCATCTGATCTTCCGACGTTAACACCACCAGGC
ACGAGCATTCTTTTTCAAGAAGCATTTTCATATGCGCTTCCAGCGCATCCCGGCGATTT
(SEQ ID NO:10)

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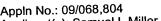
MHYFFIIVIWLLSINTAWADSGFRLKKCSILNPNYFTLSPSRNRRZNLAPLVITEMVQPI LAZCKLTASIZKGZKKWGLVKNSCYRTPAFLSLWATSILSDMMKIYGYSWEAVGAYNAGT SPKRSDIRKRYAKKIWENYRKLKGMSAEEKNKRLSIAANK

SEQ ID NO:11

WPGTICGQQHSINQQTQVKLSDGMPVPVIRLTFDGKPVALAGIRTQKIRPDRLEAHMKML LEKECSCLVVLTSERSDAGKTITTYFRGSYTFGEVHTNSQKVSSASQGEAIDQYNMQLSC GEKRYTIPVLHVKNWPDHQPLPSTDQLEYLADRVKNSNQNGAPGRSSSDKHLPMIHCLGG VGRTGTMAAALVLKDNPHSNLEQVRADFRIHGTIECWKTPAQFVQLKAMQAQLLMTTAS SEQ ID NO:12

## FIG. 25

MRDCLNNGNPVLNVGASGLTTLPDRLPPHITTLVIPDNNLTSLPELPEGLRELEVSGNLQ LTSLPSLPQGLQKLWAYNNWLASLPTLPPGLGDLAVSNNQLTSLPEMPPALRELRVSGNN LTSLRALPSGLQKLWAYNNRLTSLPEMSPGLQELDVSHNQLTRLPQSLTGLSSAARVYLD GNPLSVRTRDRLCGHHWPFRHQDTLRYGGAFRPREARALHLAVADWLTSAREGEAAQADR WQAFGLEDNAAAFSLVLDRLRETENFKKDAGFKAQISSWLTQLAEDAALRAKTFAMATEA TSTCEDRVTHALHQMNNVQLVHNAEKGEYDNNLQGLVSTGREMFRLATLEQIAREKAGTL ALVDDVEVYLAFQNKLKESLELTSVTSEMRFFDVSGVTVSDLQAADVQVKTAENSGFSKW ILQWGPLHSVLERKVPERFNALREKQISDYEDTYRKLYDEVLKSSGLVDDTDAERTIGVS AMDSAKKEFLDGLRALVDEVLGSYLTARWRLNZ SEQ ID NO:14





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|             |            | CGGCAATCCA   | GTGCTTAACG |            | <u>AGGTCTTACC</u> |
|-------------|------------|--------------|------------|------------|-------------------|
| ATGCGTGATT  | GCCTGAATAA | ACCGCATATT   | ACAACACTGG | TTATTCCTGA | TAATAATCTG        |
| ACCTTACCAG_ | ACCGTTTACC |              | CGGGAGCTGG | AGGTCTCTGG | TAACCTACAA        |
| ACCAGCCTGC  | CGGAGTTGCC | GGAAGGACTA   | CTACAGAAGC | TGTGGGCCTA | TAATAATTGG        |
| CTGACCAGCC  | TGCCATCGCT | GCCGCAGGGA   | CTAGGGGATC | TGGCGGTCTC | TAATAACCAG        |
| CTGGCCAGCC  | TGCCGACGTT | GCCGCCAGGA   | CTACGGGAGC | TGAGGGTCTC | TGGTAACAAC        |
| CTGACCAGCC  | TGCCGGAGAT | GCCGCCAGCA   | TACAGAAGCT | GTGGGCCTAT | AATAATCGGC        |
| CTGACCAGCT  | GCGCGCGCTG | CCGTCAGGAC   | TACAGGAGCT | GGATGTCTCT | CATAACCAGC        |
| TGACCAGCCT  | GCCGGAGATG | TCGCCAGGAC   | TGTCTTCAGC | GGCACGCGTA | TATCTGGACG        |
| TGACCCGCCT  | GCCGCAAAGC | CTCACGGGTC   | GGCTCTGCGG | ACATCATTGG | CCATTCAGGC        |
| GGAATCCACT  | GTCTGTACGC | ACTCGTGACA   | TCCGTCCCCG | GGAAGCCCGG | GCACTGCACC        |
| ATCAGGATAC  | ACTTCGATAT | GGCGGGGCCT   | GGGAGGGGGA | AGCGGCCCAG | GCAGACAGAT        |
| TGGCGGTCGC  | TGACTGGCTG |              | CCGCCTTCAG | CCTGGTCCTG | GACAGACTGC        |
| GGCAGGCGTT  | CGGACTGGAA | GATAACGCCG   |            | ACAGATATCA | TCCTGGCTGA        |
| GTGAGACGGA  | AAACTTCAAA | AAAGACGCGG   |            | TGCCATGGCA | ACAGAGGCAA        |
| CACAACTGGC  | TGAAGATGCT | GCGCTGAGAG   |            |            | GTACAACTGG        |
| CATCAACCTG  | CGAGGACCGG | GTCACACATG   |            | GGGGCTGGTT | TCCACGGGGC        |
| TACATAATGC  | AGAAAAAGGG | GAATACGACA   |            |            |                   |
| GTGAGATGTT  |            | ACACTGGAAC   |            |            |                   |
| CTTTAGTCGA  |            | GTCTATCTGG   |            |            |                   |
| AGCTGACCAG  |            | GAAATGCGTI   |            |            |                   |
| ACCTTCAGGC  |            | CAGGTGAAAA   |            |            |                   |
| TACTGCAGTC  |            | CACAGCGTGC   |            |            |                   |
| CGCTTCGTGA  |            | TCGGATTATC   |            |            |                   |
| TGCTGAAATC  |            | GTCGACGATA   |            |            |                   |
| CGATGGATAC  |            | A GAATTTCTGO |            | COCICIIGIO | , 0.110.00.00     |
| TGGGTAGCT   |            |              | TTAACTGA   | G          | EQ ID NO:13       |
| 10001110011 |            |              |            |            | ₩¥                |



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<u>CGCAAAGCCGAGGAAACGAACCGCATTATGGGATGTATCGGGAAAGTCCTCGGCGCGCTG</u> CTAACCATTGTCAGCGTTGTGGCCGCTGTTTTTACCGGTGGGGCGAGTCTGGCGCTGGCT GCGGTGGGACTTGCGGTAATGGTGGCCGATGAAATTGTGAAGGCGGCGACGGGAGTGTCG TTTATTCAGCAGGCGCTAAACCCGATTATGGAGCATGTGCTGAAGCCGTTAATGGAGCTG ATTGGCAAGGCGATTACCAAAGCGCTGGAAGGATTAGGCGTCGATAAGAAAACGGCAGAG ATGGCCGGCAGCATTGTTGGTGCGATTGTCGCCGCTATTGCCATGGTGGCGGTCATTGTG GTGGTCGCAGTTGTCGGGAAAGGCGCGGCGGCGAAACTGGGTAACGCGCTGAGCAAAATG ATGGGCGAAACGATTAAGAAGTTGGTGCCTAACGTGCTGAAACAGTTGGCGCAAAACGGC AGCAAACTCTTTACCCAGGGGATGCAACGTATTACTAGCGGTCTGGGTAATGTGGGTA GTGGCGTTGGGCATGGAAGTCACGAATACCGCAGCCCAGTCAGCCGGTGGTGTTGCCGAG GGCGTATTTATTAAAAATGCCAGCGAGGCGCTTGCTGATTTTATGCTCGCCCGTTTTGCC ATGGATCAGATTCAGCAGTGGCTTAAACAATCCGTAGAAATATTTGGTGAAAACCAGAAG GTAACGGCGGAACTGCAAAAAGCCATGTCTTCTGCGGTACAGCAAAATGCGGATGCTTCG <u>CGTTTTATTCTGCGCCAGAGTCGCGCATAA</u>AAACTGCCAAAATAAAGGGAGAAAAAT<mark>ATG</mark> TTAATTAGTAATGTGGGAATAAATCCCGCCGCTTATTTAAATAATCATTCTGTTGAGAAT <u>AGTTCACAGACAGCTTCGCAATCCGTTAGCGCTAAAGATATTCTGAATAGTATTGGTATT</u> AGCAGCAGTAAAGTCAGTGACCTGGGGTTGAGTCCTACACTGAGCGCCCTGCGCCAGGG GTATTAACGCAAACCCCCGGAACGATCACGTCCTTTTTAAAAGCCAGTATTCAAAATACC GACATGAATCAGGATTTGAATGCTCTGGCAAATAATGTCACGACTAAAGCGAATGAGGTT GTGCAAACCCAGTTACGCGAGCAGCAGGCAGAAGTCGGAAAGTTTTTTGATATTAGCGGA ATGTCTTCCAGTGCCGTTGCGCTGTTGGCTGCCGCGAATACGTTAATGCTGACGTTGAAC CAGGCTGATAGCAAACTGTCTGGTAAGTTGTCATTAGTCAGTTTTGATGCAGCTAAAACG ACGGCAAGCTCCATGATGCGCGAAGGGATGAATGCGTTGTCCGGTAGTATTTCCCAGAGC GCGCTTCAGTTGGGGATCACTGGCGTGGGCGCCAAACTGGAATATAAGGGGCTGCAGAAT GAAAGAGGCGCGCTTAAACATAATGCCGCGAAGATCGATAAACTGACCACTGAAAGCCAC <u>AGTATTAAAAACGTGCTGAACGGGCAGAATAGCGTCAAACTCGGTGCTGAAGGCGTCGAT</u> TCTCTGAAATCGTTAAATATGAAGAAAACCGGTACCGATGCGACGAAAAATCTTAATGAT GCGACGCTTAAATCTAATGCCGGAACCAGCGCCACGGAAAGTCTGGGTATTAAAGACAGT <u>AATAAACAAATCTCCCCTGAACATCAGGCTATTCTGTCGAAACGTCTTGAGTCTGTCGAA</u> TCCGATATTCGTCTTGAGCAGAATACCATGGATATGACCCGAATCGATGCGCGCAAGATG TCCGGGCAGTACGCCGCTACTCAGGAACGTTCCGAGCAGCAAATTAGCCAGGTGAATAAC CGGGTTGCCAGCACCGCATCGGACGAAGCCCGTGAAAGTTCACGTAAATCGACCAGCCTG <u>ATTCAGGAAATGCTGAAAACAATGGAGAGCATTAACCAGTCGAAAGCATCCGCACTCGCT</u> GCTATCGCAGGCAATATTCGCGCTTAATCTGAAAGGTCATCTATACGCCATCATGGGTGT GATTTAATCGCGCTCCTGATGGCGAACTGGGGATATT<u>ATGCTTAATATTCAAAATTA</u> CGTCGAGACTGCCGTGGTACCGTCTACCACAGAACATCGCGGTACAGATATCATTTCATT ATCGCAGGCGGCTACTAAAATCCACCAGGCACAGCAGACGCTGCAGTCAACGCCACCGAT CTCTGAAGAGAATAATGACGAGCGCACGCTGGCGCCCAGCAGTTGACCAGCAGCCTGAA TGCGCTGGCGAAGTCCGGCGTGTCATTATCCGCAGAACAAAATGAGAACCTGCGGAGCGC <u>GTTTTCTGCGCCGACGTCGGCCTTATTTAGCGCTTCGCCTATGGCGCAGCCGAGAACAAC</u> CATTTCTGATGCTGAGATTTGGGATATGGTTTCCCAAAATATATCGGCGATAGGTGACAG <u>CTATCTGGGCGTTTATGAAAACGTTGTCGCAGTCTATACCGATTTTTATCAGGCCTTCAG</u> <u>TGATATTCTTTCCAAAATGGGAGGCTGGTTATTACCAGGTAAGGACGGTAATACCGTTAA</u> AAACAGTAATACCGTTTTATTTCCAGCGCAGTCAGGCAGCGGCGTTAAAGTAGCCACTGA <u>AGCGGAAGCGAGACAGTGGCTCAGTGAATTGAATTTACCGAATAGCTGCCTGAAATCTTA</u> TGGATCCGGTTATGTCGTCACCGTTGATCTGACGCCATTACAAAAAATGGTTCAGGATAT TGATGGTTTAGGCGCGCGGGAAAAGACTCAAAACTCGAAATGGATAACGCCAAATATCA AGCCTGGCAGTCGGGTTTTAAAGCGCAGGAAGAAAATATGAAAACCACATTACAGACGCT GACGCAAAAATATAGCAATGCCAATTCATTGTACGACAACCTGGTAAAAGTGCTGAGCAG <u>TACGATAAGTAGCAGCCTGGAAACCGCCAAAAGCTTCCTGCAAGGATAA</u>CAGAAGAGGAT ATTAATAATGGTTACAAGTGTAAGGACTCAGCCCCCGTCATAATGCCAGGTATGCAGAC CGAGATCAAAACGCAGGCCACGAATCTTGCGGCGAATCTTTCCGCAGTCAGAGAAAGTGC CACAGCGACGCTGTCAGGGGAAATTAAAGGCCCGCAACTGGAAGATTTTCCCGCGCTGAT <u>CAAACAGGCGAGTCTGGATGC</u> SEQ ID NO:15



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GAGCTCAGCAACGTGTCGAAAGCCTGTAAAATCATGGGCGTCTCGCGCGATACGTTTTAC CGTTATCGTGAACTGGCCGATGAAGGCGGCGTTGATGCGCTGATAAATCGTAGTCGCCGC GTACCTAACCTTAAGAACCGTACCGATGAGGCAACTGAGCAAGCTGTTGTTGATTATGCC GTTGCGTTCCCGGCCCATGGTCAGCACCGAACTGCGCAAACAGGACGTTTTTATCTCCGG TAGTGATGTCCATTCCGTCTGGCTGCGCACAACCTTGAGAACTTCAAAAAACGCCTGAAA AGCGTAAAGCCAGTGATGATGAAGCCTGTGGTGAGATTGAAACCGTTCATCCGGGATATC TGGGGTCACAGGACACGTTCTACGTGGGCAACCTGAAAGGCGTTGGGCGAATCTATCAGC AGACGTTCGTTGATACATACTCGAAGGTGGCTCACTGCAAGCGCTATATCACCAAAACGC CGATTACAGCGGCTGATTTGCTGAATGATCGTGTACTGCCGTTTATGAGTCTCAGGGCCT GCCGATGCTAAGGATACTGACAGACAGGGGTACAGAATATTGCGGCAAAGTGGAACATCA TGATTATCAGCTTTATCTGGAGATAAATGACATCGAACACACGAAAACGAAGGCGATGTC CCCGCAGACCAATGGCATCTGCGAGCGGTTCCATAAAACGATACTGAACGAATTTTATCA GGTGACGTTCCGCAAAAAGTTATATGGCGATTTTGATACATTACAATCGGATCTTGATGA ATGGCTGGTTCACTATAATAATGAGCGAACCCATCAGGGAAAAATGTGCTGTGGCCGGAC GCCGATGGAAACGTTACTTGATGGAAAACGCATCTGGTCTGAGAAGAATTTAAGCCAGAT GTAATCTGACAGATACCTGTATAAATAACCGGTAACTGTCAGATCAGGTCTGAGCTAATA ATGCGTGTGTCTGCGTTAATCTCTTTTCATTGTGCTGTAAATTAGGCAGTGGAATATGTT TAATATCCGCAATACACAACCTTCTGTAAGTATGCAGGCTATTGCTGGTGCAGCGGCACC AGAGGCATCTCCGGAAGAAATTGTATGGGAAAAATTCAGGTTTTTTTCCCGCAGGAAAAT TACGAAGAAGCGCAACAGTGTCTCGCTGAACTTTGCCATCCGGCCCGGGGAATGTTGCCT GATCATATCAGCAGCCAGTTTGCGCGTTTAAAAGCGCTTACCTTCCCCGCGTGGGAGGAG GAGATATTGTCAATCACTCTTGATGATGCCGGGAACTATACCGTGAATTGTCAGGGGTAC AGTGAAGCACATGACTTCATCATGGACACAGAACCGGGAGAGGAATGCACAGAATTCGCG GAGGGGGCATCCGGGACATCCCTCCGCCCTGCCACAACGGTTTCACAGAAGGCAGCAGAG TATGATGCTGTCTGGTCAAATGGGAAAGGGATGCACCAGCAGGAGAGTCACCCGGCCGCG CAGCAGTGGTACAGGAAATGCGTGATTGCCTGAATAACGGCAATCCAGTGCTTAACGTGG **GAGCGTCAGGTCTTACCACCTTACCAGACCGTTTACCACCGCATATTACAACACTGGTTA** <u>TTCCTGATAATAATCTGACCAGCCTGCCGGAGTTGCCGGAAGGACTACGGGAGCTGGAGG</u> <u>TCTCTGGTAACCTACAACTGACCAGCCTGCCATCGCTGCCGCAGGGACTACAGAAGCTGT</u> <u>GGGCCTATAATAATTGGCTGGCCAGCCTGCCGACGTTGCCGCCAGGACTAGGGGATCTGG</u> CGGTCTCTAATAACCAGCTGACCAGCCTGCCGGAGATGCCGCCAGCACTACGGGAGCTGA <u>GGGTCTCTGGTAACAACCTGACCAGCTGCGCGCGCTGCCGTCAGGACTACAGAAGCTGTG</u> GGCCTATAATAATCGGCTGACCAGCCTGCCGGAGATGTCGCCAGGACTACAGGAGCTGGA <u>ACGCGTATATCTGGACGGGAATCCACTGTCTGTACGCACTCGTGACAGGCTCTGCGGACA</u> <u>TCATTGGCCATTCAGGCATCAGGATACACTTCGATATGGCGGGGCCTTCCGTCCCCGGGA</u> <u>AGCCCGGGCACTGCACCTGGCGGTCGCTGACTGGCTGACGTCTGCACGGGAGGGGGAAGC</u> <u>GGCCCAGGCAGACAGATGGCAGGCGTTCGGACTGGAAGATAACGCCGCCGCCTTCAGCCT</u> <u>GGTCCTGGACAGACTGCGTGAGACGGAAAACTTCAAAAAAGACGCGGGCTTTAAGGCACA</u> <u>GATATCATCCTGGCTGACACAACTGGCTGAAGATGCTGCGCTGAGAGCAAAAACCT</u> TTGCCATGGCAACAGAGGCAACATCAACCTGCGAGGACCGGGTCACACATGCCCTGCACC <u>AGATGAATAACGTACAACTGGTACATAATGCAGAAAAAGGGGAATACGACAACAATCTCC</u> <u>AGGGGCTGGTTTCCACGGGGCGTGAGATGTTCCGCCTGGCAACACTGGAACAGATTGCCC</u> <u>GGGAAAAAGCCGGAACACTGGCTTTAGTCGATGACGTTGAGGTCTATCTGGCGTTCCAGA</u> <u>ATAAGCTGAAGGAATCACTTGAGCTGACCAGCGTGACGTCAGAAATGCGTTTCTTTGACG</u> TTTCCGGCGTGACGGTTTCA GACCTTCAGGCTGCGGACGTTCAGGTGAAAACCGCTGAAAACAGCGGGTTCAGTAAATGG <u>ATACTGCAGTGGGGGCCGTTACACAGCGTGCTGGAACGCAAAGTGCCGGAACGCTTTAAC</u> <u>GCGCTTCGTGAAAAGCAAATATCGGATTATGAAGACACGTACCGGAAGCTGTATGACGAA</u> <u>GTGCTGAAATCGTCCGGGCTGGTCGACGATACCGATGCAGAACGTACTATCGGAGTAAGT</u> <u>GCGATGGATAGTGCGAAAAAAGAATTTCTGGATGGCCTGCGCGCTCTTGTGGATGAGGTG</u> CTGGGTAGCTATCTGACAGCCCGGTGGCGTCTTAACTGAGCACGATATTCTCCGCACCAG GATGCACAGGTGAAACAGGGGAGACTTCTTCAGTCAGGGCGTACGCAACTCAACCTTTTC GACGATACGCGCC/ SEQ ID NO:16

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